Network Systems
Science & Advanced
Computing

Biocomplexity Institute & Initiative

University of Virginia

Estimation of COVID-19 Impact in Virginia

October 13th, 2021

(data current to October 9th – 12th)
Biocomplexity Institute Technical report: TR 2021-110



BIOCOMPLEXITY INSTITUTE

biocomplexity.virginia.edu

About Us

- Biocomplexity Institute at the University of Virginia
 - Using big data and simulations to understand massively interactive systems and solve societal problems
- Over 20 years of crafting and analyzing infectious disease models
 - Pandemic response for Influenza, Ebola, Zika, and others



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Overview

• Goal: Understand impact of COVID-19 mitigations in Virginia

Approach:

- Calibrate explanatory mechanistic model to observed cases
- Project based on scenarios for next 4 months
- Consider a range of possible mitigation effects in "what-if" scenarios

Outcomes:

- Ill, Confirmed, Hospitalized, ICU, Ventilated, Death
- Geographic spread over time, case counts, healthcare burdens

Key Takeaways

Projecting future cases precisely is impossible and unnecessary. Even without perfect projections, we can confidently draw conclusions:

- Case rates in Virginia continue to decline, with nearly all districts declining as well; case rates remain high though the rate of decline has picked up again
- VA 7-day mean daily incidence is slightly down to 29/100K from 32/100K; US is also slightly down to 35/100K (from 39/100K)
- Projections show continued decline across the board, more rapid in some districts than others
- Future case growth remains possible, however, when tested with transmission drivers from last year
- Recent updates:
 - Analysis to show potential impact of Influenza based on past seasons
 - Updated vaccination schedules, adjusted 3rd dose uptake and 5-11 year old timing
 - Adjustment to higher levels of assumed immunity waning (natural and vaccine)

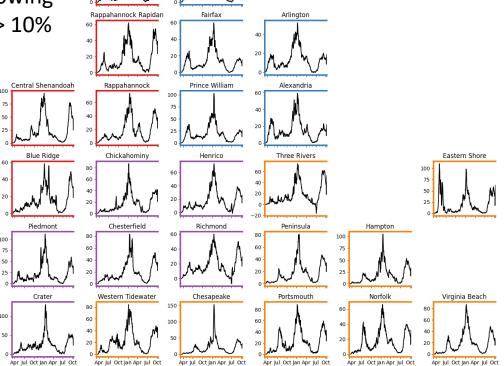
The situation continues to change. Models continue to be updated regularly.

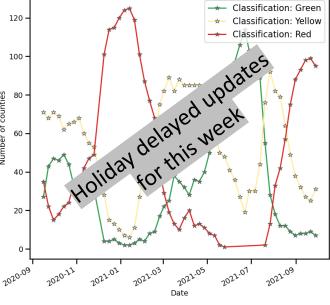
Situation Assessment



Case Rates (per 100k) and Test Positivity

- Case rate increase across all health districts
- Some past 50% of winter peak and growing
- More than 50% of counties with TPR > 10%





County level RT-PCR test positivity

Green: <5.0% (or <20 tests in past 14 days)
Yellow: 5.0%-10.0% (or <500 tests and <2000
tests/100k and >10% positivity over 14 days)
Red: >10.0% (and not "Green" or "Yellow")

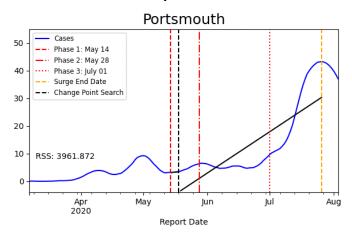
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District Trajectories

Goal: Define epochs of a Health District's COVID-19 incidence to characterize the current trajectory

Method: Find recent peak and use hockey stick fit to find inflection point afterwards, then use this period's slope to define the trajectory

Hockey stick fit



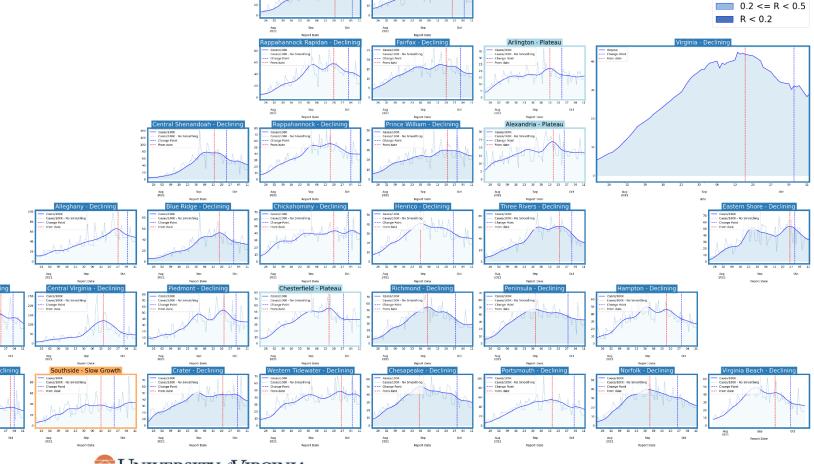
Trajectory	Description	Weekly Case Rate (per 100K) bounds	# Districts (prev week)
Declining	Sustained decreases following a recent peak	below -0.9	31 (31)
Plateau	Steady level with minimal trend up or down	above -0.9 and below 0.5	3 (3)
Slow Growth	Sustained growth not rapid enough to be considered a Surge	above 0.5 and below 2.5	1 (1)
In Surge	Currently experiencing sustained rapid and significant growth	2.5 or greater	0 (0)



District Trajectories – last 10 weeks

Status	# Districts (prev week)
Declining	31 (31)
Plateau	3 (3)
Slow Growth	1 (1)
In Surge	0 (0)

Curve shows smoothed case rate (per 100K)
Trajectories of states in label & chart box
Case Rate curve colored by Reproductive
number



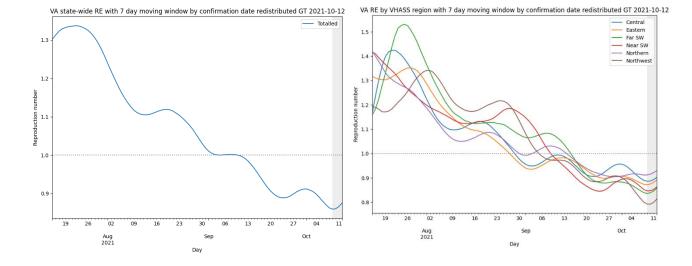


Estimating Daily Reproductive Number –

Redistributed gap

Oct 12th Estimates

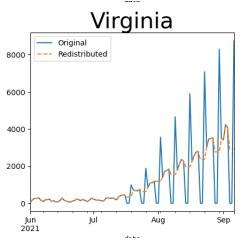
Region	Date Confirmed R _e	Date Confirmed Diff Last Week
State-wide	0.889	-0.125
Central	0.902	-0.139
Eastern	0.892	-0.115
Far SW	0.858	-0.130
Near SW	0.863	-0.121
Northern	0.930	-0.059
Northwest	0.813	-0.206



Skipping Weekend Reports & holidays biases estimates
Redistributed "big" report day to fill in gaps, and then estimate R from
"smoothed" time series

Methodology

- Wallinga-Teunis method (EpiEstim¹) for cases by confirmation date
- Serial interval: updated to discrete distribution from observations (mean=4.3, Flaxman et al, Nature 2020)
- Using Confirmation date since due to increasingly unstable estimates from onset date due to backfill

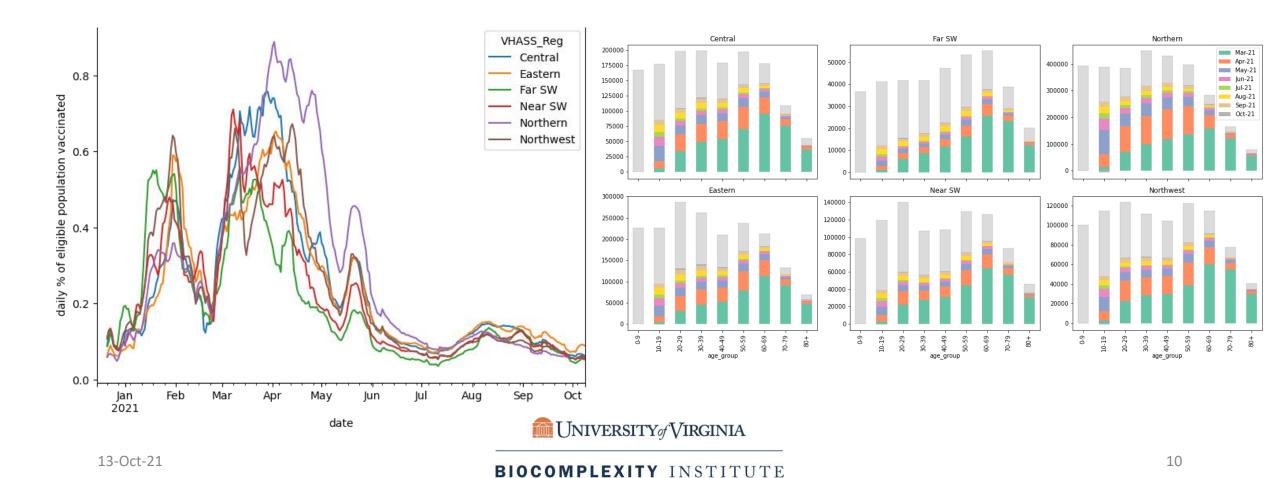


^{1.} Anne Cori, Neil M. Ferguson, Christophe Fraser, Simon Cauchemez. A New Framework and Software to Estimate Time-Varying Reproduction Numbers During Epidemics. American Journal of Epidemiology, Volume 178, Issue 9, 1 November 2013, Pages 1505–1512, https://doi.org/10.1093/aje/kwt133

Vaccination Administration Slow

Regional Vaccine courses initiated per day (% eligible):

- Proportion eligible for first dose of vaccines across regions (in the ~0.1% or 100 per 100K a day)
- Age-specific proportions of population vaccinated show recent progress in younger ages

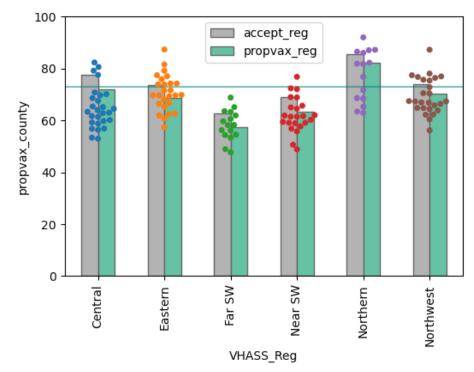


Vaccination Acceptance by Region

Corrections to surveys:

- Facebook administered survey is timely and broad, but biased by who accesses Facebook and answers the survey
- Correction approach:
 - Calculate an over-reporting fraction based on reported vaccinations compared to VDH administration data
 - Cross-validate coarse corrections against HPS survey at the state level and corrected in same manner

Region	COVIDcast accepting corrected	VDH proportion pop vaccinated
Central	77%	72%
Eastern	74%	68%
Far SW	63%	57%
Near SW	71%	63%
Northern	85%	82%
Northwest	75%	70%
Virginia	78%	72%



Grey Bar: Survey measured and corrected acceptance

Green Bar: Proportion of eligible population

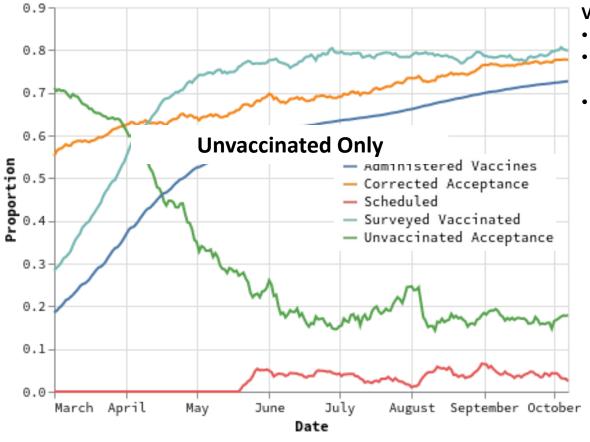
administered a vaccine

Dots: Proportion administered at least one dose for

each county



Vaccine Acceptance Components over Time

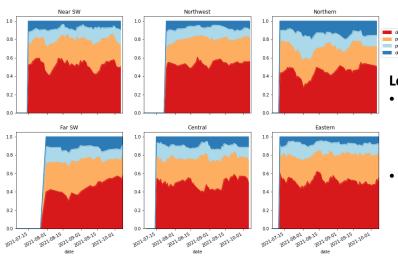


Vaccine Acceptance adjusted to include scheduled appointments

- Steady rise in acceptance over the past couple months
- Unvaccinated Acceptance shows ~20% of those who are unvaccinated are definitely or probably willing to be vaccinated
- Scheduled appointments for vaccination have increased through August but seem to be leveling off

Levels of Vaccine Willingness





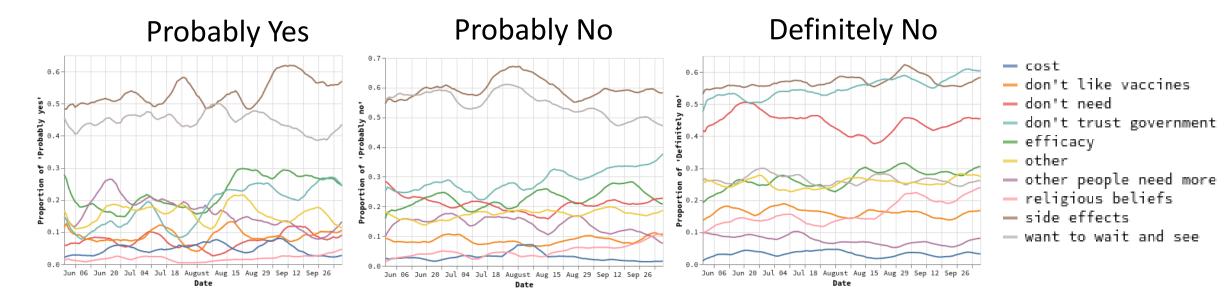
Levels of Acceptance in flux:

- Most regions are steady with 20-30% of unvaccinated still in the Definitely/Probably "Yes" categories.
- About 50% of the Unvaccinated seem to be in the "Definitely Not" category.

Data Source: https://covidcast.cmu.edu



Reasons for Hesitancy by Likeliness to Accept



Reasons for Hesitancy vary across tiers of likeliness to accept the vaccine

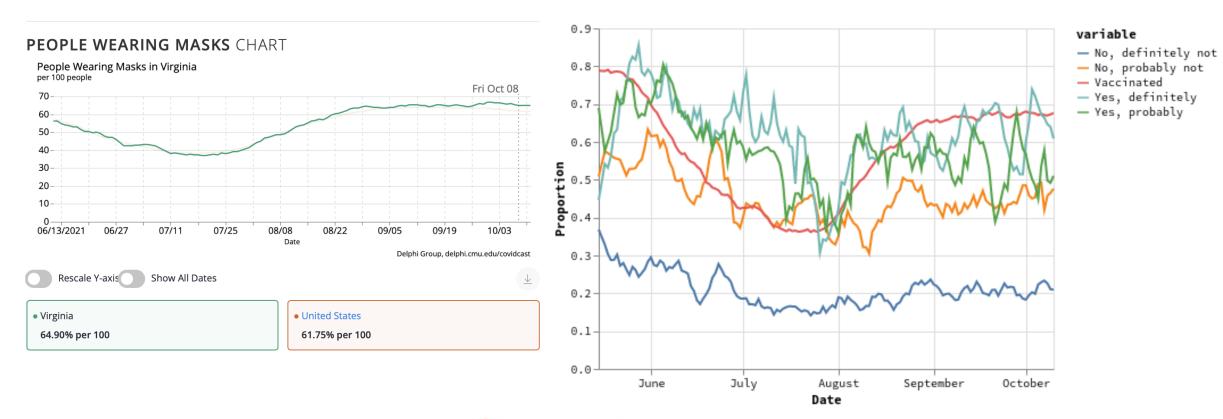
- Probably Yes and Probably No most concerned about side effects & are waiting to see
- Definitely No are concerned about side effects but also don't think they need the vaccine and don't trust the government, though don't need is declining
- Most other reasons are below 30% within these tiers of likeliness



Mask Usage Stalls

Self-reported mask usage has plateaued out to ~65%, perhaps slight decline starting

- US and VA similar, though with considerable variation across counties and states
- Mask wearing remains lower amongst unvaccinated especially among least willing to be vaccinated



Data Source: https://covidcast.cmu.edu



SARS-CoV2 Variants of Concern

Emerging new variants will alter the future trajectories of pandemic and have implications for future control

- Emerging variants can:
 - · Increase transmissibility
 - Increase severity (more hospitalizations and/or deaths)
 - Limit immunity provided by prior infection and vaccinations
- Genomic surveillance remains very limited
 - Challenges ability to estimate impact in US to date and estimation of arrival and potential impact in future

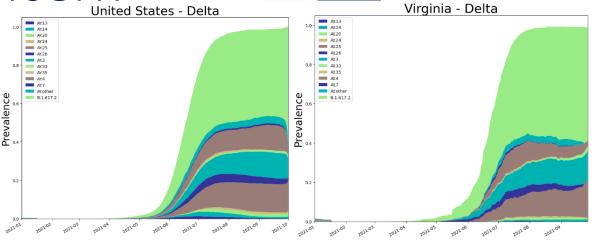
	New WHO Name	Transmissibility	Immune Evasiveness	Vaccine Effectiveness^
Ancestral		_	_	✓
D614G		+		✓
B.1.1.7	Alpha	+++	· <u></u> -	✓
B.1.351	Beta	+	++++	✓
P.1	Gamma	++	++	✓
B.1.429	Epsilon	+	+	✓
B.1.526	lota	+	+	✓
B.1.617.2	Delta	++++*	++#	✓

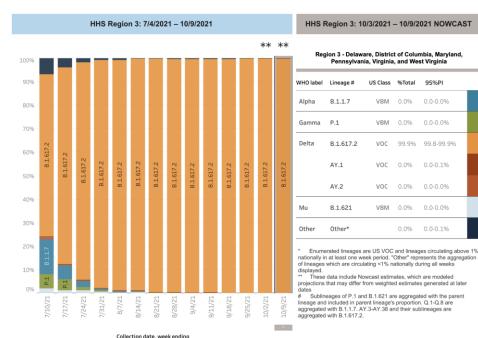
^{*}Relative transmissibility to B.1.1.7 yet to be fully defined

[^]Effectiveness from real world evidence vs. severe illness, not all vaccines are effective vs all variants, and importance of 2-doses, especially for B.1.617.2 for which 1 dose of mRNA or AZ is only ~30% effective # May carry more immune escape than P.1, to be determined



WHO and Eric Topol

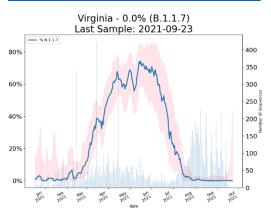




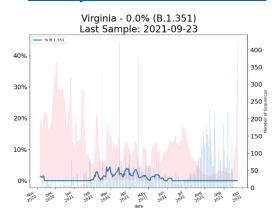
SARS-CoV2 Variants of Concern

Previous Variants

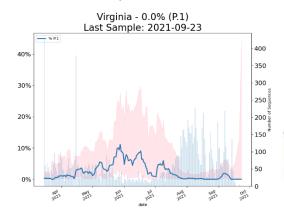
Alpha α - Lineage B.1.1.7

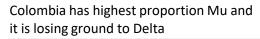


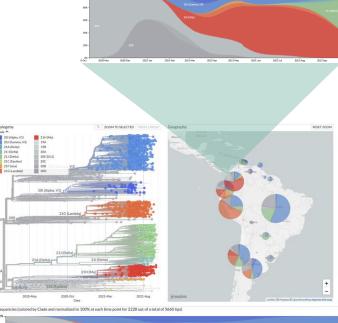
Beta β - Lineage B.1.351

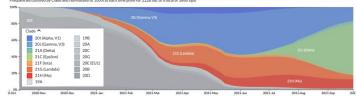


Gamma γ - **Lineage P.1**





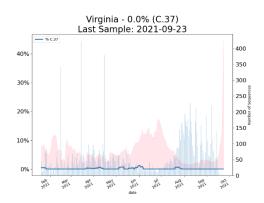




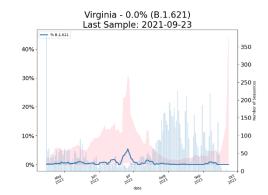
Delta continues to out compete Lambda and Mu in South America (over 60%, Lambda and Mu both below 10%)

<u>Trevor Bedford Tweet & Nextstrain Analysis</u>

Emerging Variants Lambda λ - **Lineage C.37**



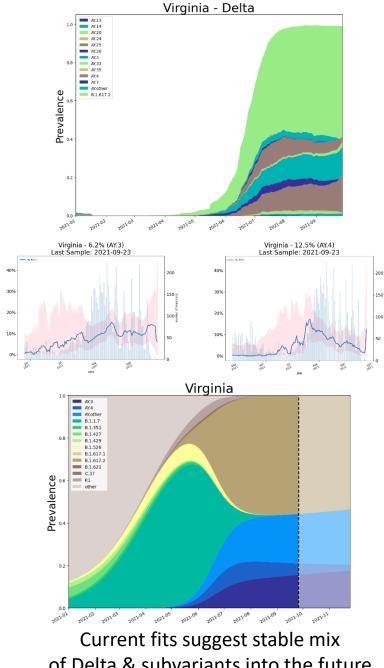
Mu μ - **Lineage B.1.621**



SARS-CoV2 Variants of Concern

Delta δ - **Lineage B.1.617.2** and related subvariants

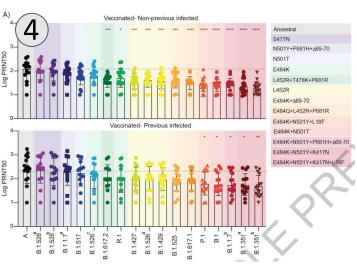
- Delta plus δ + lineage which contains the K417N mutation is emerging as a sub-variant that is even more transmissible; declared a VoC in India
- Delta variant now dominates most of Europe and US
- CDC recommends resumption of mask wearing indoors due to reports of breakthrough infections of the vaccinated possibly being transmissible
- Recent study from Mayo clinic shows Delta reducing the efficacy of mRNA vaccines (Pfizer more so than Moderna) along with other reports. Israeli study showed 64% efficacy against infection, however, a 3rd dose may counteract this reduction
- Public Health Scotland study in Lancet suggests Delta is 2x more likely to cause hospitalization than Alpha
- Subvariants AY.3 and AY.4 with some significant variability while there are limited genomes submitted, these subvariants are mainly clustered in the US, others mainly outside of US



of Delta & subvariants into the future

Variants & Vaccines

- Partial or dense exposure to viral particles implicated in severity. These results reinforce the importance of public health measures that limit exposure dose, such as social distancing, masking, and increased ventilation.
- Potentially largest study of a third dose of BNT162b2 vaccine in health-care workers shows high vaccine immunoreactivity. The median time between the first and third vaccine doses was 32.0 weeks. Isreali cohorts such as this one provide valuable data on robust immunity of large interval vaccination schedules.
- Results demonstrate that the concept of vaccine bubbles is valuable, especially as we approach the holiday season.
- "Plasma from previously infected vaccinated individuals displayed overall better neutralization capacity when compared to plasma from uninfected individuals that also received two vaccine doses, pointing to vaccine boosters as a relevant future strategy to alleviate the impact of emerging variants on antibody neutralizing activity."



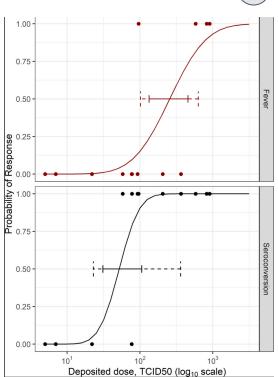
Recent Yale study demonstrates mRNA vaccines induce robust antibodies and T-cells; certain mutations increase immune escape; prior infection + 2 doses produces very high neutralizing concentrations against most variants

https://www.nature.com/articles/s41586-021-04085-y reference.pdf

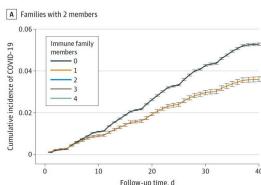


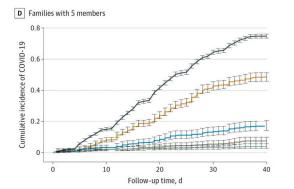
National Biodefense Analysis and Countermeasures Center (NBACC) researchers conduct a study on 16 macaques and are the first to demonstrate that the probability of infection is dependent on the exposure dose of SARS-CoV-2 in a nonhuman primate model of inhalational COVID-19. The results demonstrate that the probability of seroconversion and fever are both dose-dependent, but that the median dose for seroconversion is significantly lower than that of fever, resulting in a group of animals that developed an immune response post-exposure but did not develop fever or other clinical signs of infection

https://journals.plos.org/plospathog ens/article?id=10.1371/journal.ppat. 1009865



Cohort study of 1,789,728 individuals from 814,806 families in Sweden, family members without immunity had a 45% to 97% lower risk of contracting COVID-19 as the number of immune family members increased. https://jamanetwork.com/journals/jamainternalmedicine/fullarticle/2785141





0.5 Before Third Dose After Third Dose A new Isreali study demonstrates substantial IgG response of 346 healthcare workers (median age 67) induced by a 3rd shot/booster Pfizer vaccine. sCOVG index of serology assay has considered positive for SARS-CoV-2 antibodies.

50-

sCOVG Index

n=331

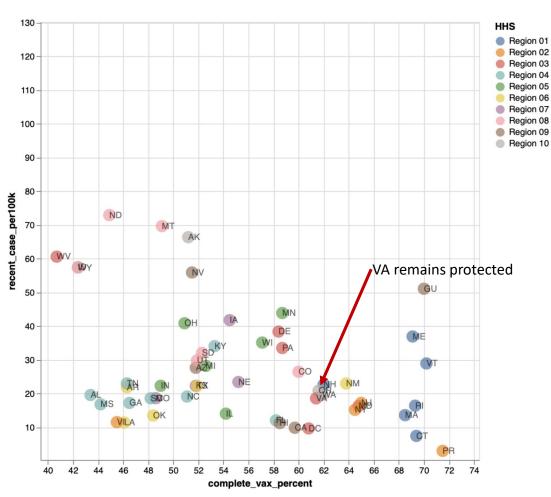
values up to 150 (upper limit of quantification), 1.00 or greater is

https://www.thelancet.com/journals/lanmic/article/PIIS2666-5247(21)00272-X/fulltext

Recent Cases Correlate with Vax Coverage

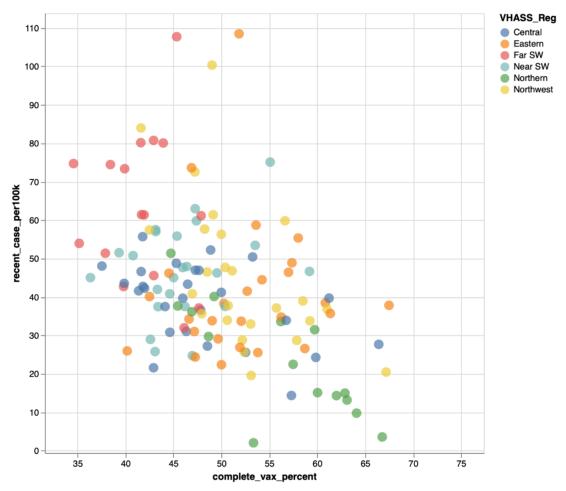
Mean cases per 100K vs. vaccine coverage

• States with lower vax coverage have had the worst case spikes

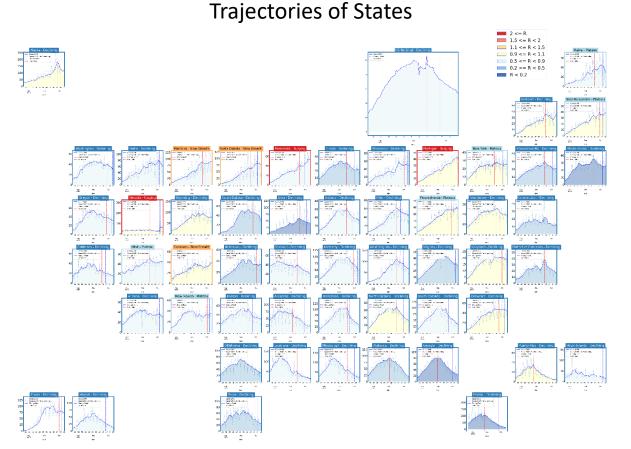


Virginia Counties

Counties with higher vax coverage are maintaining lower case rates

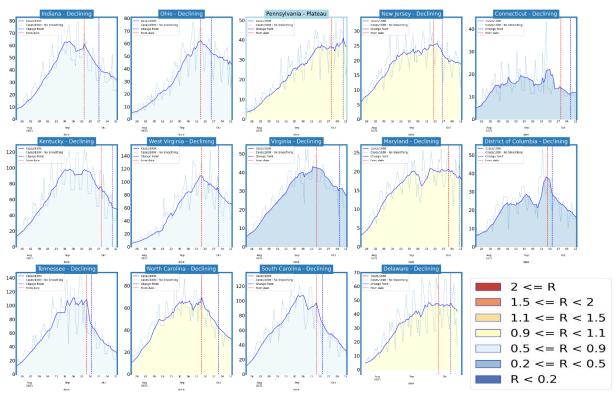


Other State Comparisons



- Most states continue to decline
- Some states remain in growth, but show signs of slowing
- Case rates remain very high, but nationally rates have been in sustained decline

Virginia and her neighbors

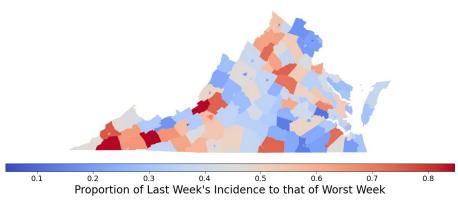


- Nearly all states now in sustained decline in for several weeks or are exiting a prolonged plateau
- Case rates remain high

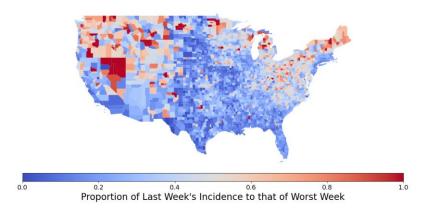
Last Week compared to the worst and the best

County level Case Rates (per 100K) proportion when comparing this most recent week to:

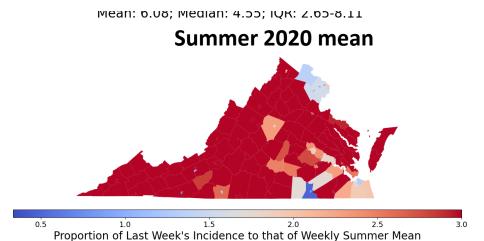
Worst Week of the Pandemic



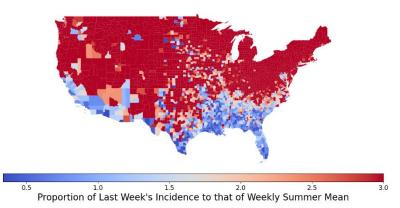
Recent Incidence Compared to Worst Week by County







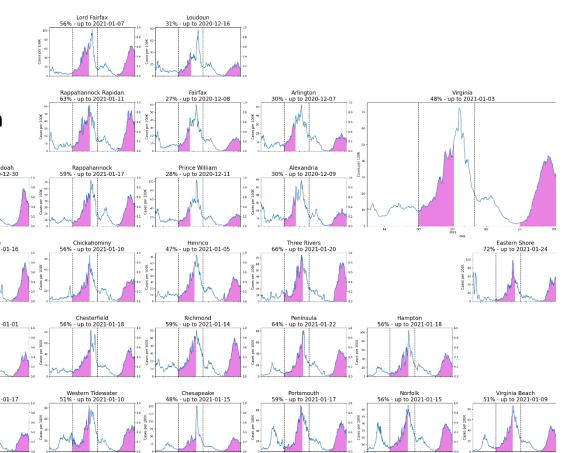
Recent Incidence Compared to Weekly Summer Mean by County Mean: 61.42: Median: 4.11: IOR: 1.8-9.69



Delta Wave compared to Last Fall – Winter wave

Total cases in Delta wave compared to cases in Fall wave and progress

- Virginia's Delta wave has caused 48% of the total cases experienced in the Fall-Winter wave
- Total for this wave to date is similar to what had been experienced by Jan 3rd in Fall-Winter wave
- Progress against the Winter wave varies by district, ranging from ~25% to over 60%

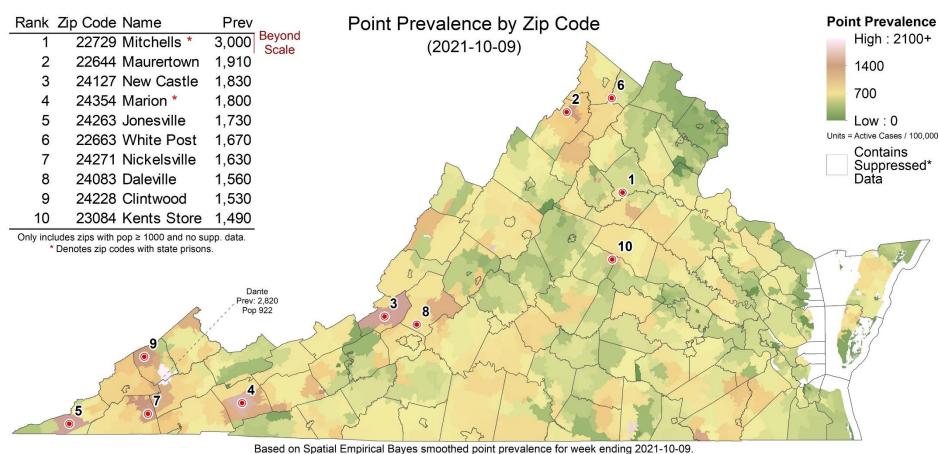


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Zip code level weekly Case Rate (per 100K)

Case Rates in the last week by zip code

- Color scaled adjusted to accommodate the very high prevalence levels this week
- Clusters of high prevalence in Southwest and Northwest
- Some counts are low and suppressed to protect anonymity, those are shown in white

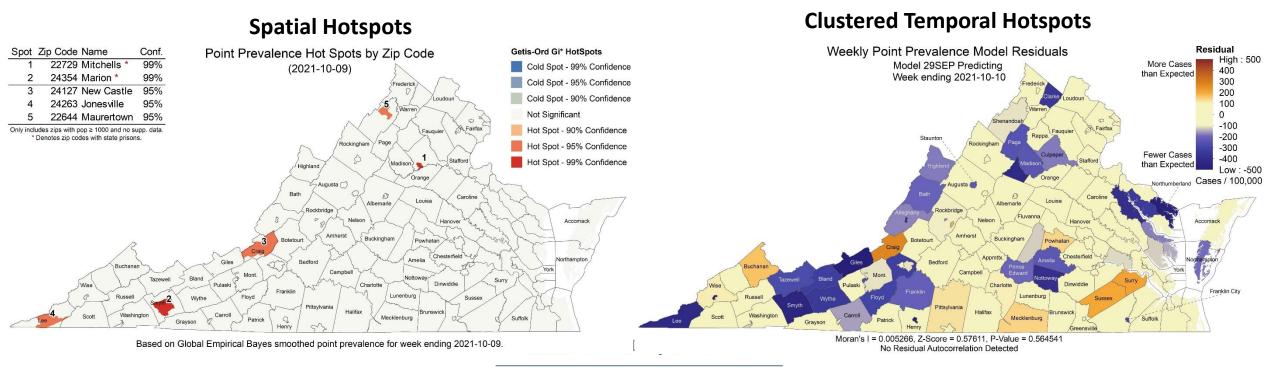




Current Hot-Spots

Case rates that are significantly different from neighboring areas or model projections

- **Spatial**: Getis-Ord Gi* based hot spots compare clusters of zip codes with weekly case prevalence higher than nearby zip codes to identify larger areas with statistically significant deviations
- **Temporal**: The weekly case rate (per 100K) projected last week compared to observed by county, which highlights temporal fluctuations that differ from the model's projections



Model Update – Adaptive Fitting



Adaptive Fitting Approach

Each county fit precisely, with recent trends used for future projection

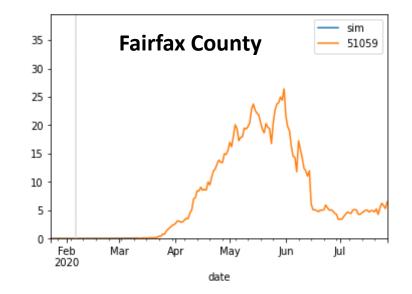
 Allows history to be precisely captured, and used to guide bounds on projections

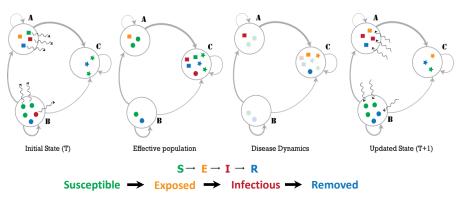
Model: An alternative use of the same meta-population model, PatchSim

- Allows for future "what-if" Scenarios to be layered on top of calibrated model
- Eliminates connectivity between patches, to allow calibration to capture the increasingly unsynchronized epidemic

External Seeding: Steady low-level importation

- Widespread pandemic eliminates sensitivity to initial conditions
- Uses steady 1 case per 10M population per day external seeding







Using Ensemble Model to Guide Projections

Ensemble methodology that combines the Adaptive with machine learning and statistical models such as:

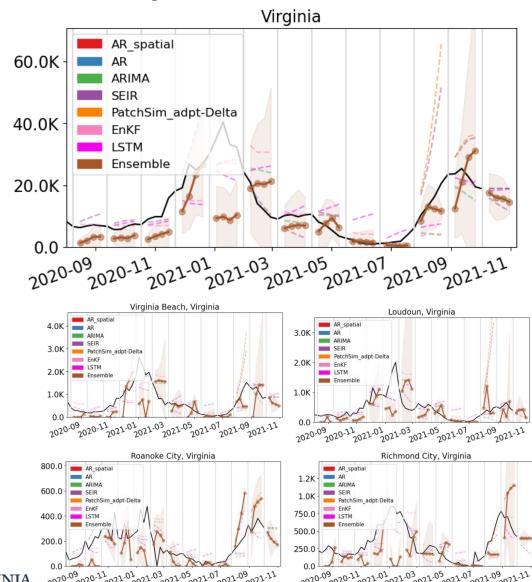
- Autoregressive (AR, ARIMA)
- Neural networks (LSTM)
- Kalman filtering (EnKF)

Weekly forecasts done at county level.

Models chosen because of their track record in disease forecasting and to increase diversity and robustness.

Ensemble forecast provides additional 'surveillance' for making scenario-based projections.

Also submitted to CDC Forecast Hub.



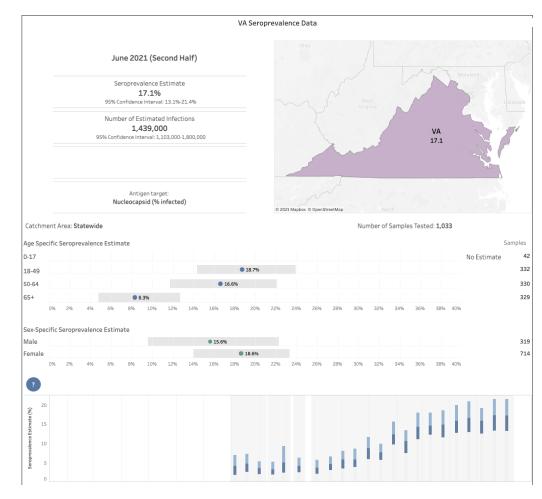
Seroprevalence updates to model design

Several seroprevalence studies provide better picture of how many actual infections have occurred

 CDC Nationwide Commercial Laboratory Seroprevalence Survey

These findings are equivalent to an ascertainment ratio of ~2x in the future, with bounds of (1.3x to 3x)

- Thus for 2x there are 2 total infections in the population for every confirmed case recently
- This measure now fully tracks the estimated ascertainment over time
- Uncertainty design has been shifted to these bounds (previously higher ascertainments as was consistent earlier in the pandemic were being used)



https://covid.cdc.gov/covid-data-tracker/#national-lab



Calibration Approach

- Data:
 - County level case counts by date of onset (from VDH)
 - Confirmed cases for model fitting
- Calibration: fit model to observed data and ensemble's forecast.
 - Tune transmissibility across ranges of:
 - Duration of incubation (5-9 days), infectiousness (3-7 days)
 - Undocumented case rate (1x to 7x) guided by seroprevalence studies
 - Detection delay: exposure to confirmation (4-12 days)
 - Approach captures uncertainty, but allows model to precisely track the full trajectory of the outbreak
- Project: future cases and outcomes generated using the collection of fit models run into the future
 - Mean trend from last 7 days of observed cases and first week of ensemble's forecast used
 - Outliers removed based on variances in the previous 3 weeks
 - 2 week interpolation to smooth transitions in rapidly changing trajectories
- Outcomes: Data driven by shift and ratio that has least error in last month of observations
 - Hospitalizations: 3 days from confirmation, 6.8% of cases hospitalized
 - Deaths: 11 days from confirmation, 1.45% of cases die





COVID-19 in Virginia:



Dashboard Updated: 10/13/2021 Data entered by 5:00 PM the prior day.

		Cases, Hospitaliza	itions and Deaths		
Total Cases* 897,978 (New Cases: 2,359)^		Total Hospitalizations** 37,613		Total Deaths 13,285	

^{*} Includes both people with a positive test (Confirmed), and symptomatic with a known exposure to COVID-19 (Probable).

[†] VDH adopted the updated CDC COVID-19 2021 Surveillance Case Definition on September 1, 2021 which is found here: -- https://ndc.services.cdc.gov/case-definitions/coronavirus-disease-2019-2021/

Outbreaks			
Total Outbreaks*	Outbreak Associated Cases		
4,726	86,978		

^{*} At least two (2) lab confirmed cases are required to classify an outbreak.

Testing (PCR Only)			
Testing Encounters PCR Only* Current 7-Day Positivity Rate PCR Only*			
9,588,161	7.6%		

^{*} PCR" refers to "Reverse transcriptase polymerase chain reaction laboratory testing."

^{**} Lab reports may not have been received yet. Percent positivity is not calculated for days with incomplete data

Multisystem Inflammatory Syndrome in Children			
Total Cases*	Total Deaths		
91	0		

^{*}Cases defined by CDC HAN case definition: https://emergency.cdc.gov/han/2020/han00432.asp

Accessed 10:30am October 13, 2021 https://www.vdh.virginia.gov/coronavirus/

^{**} Hospitalization of a case is captured at the time VDH performs case investigation. This underrepresents the total number of hospitalizations in Virginia.

[^]New cases represent the number of confirmed and probable cases reported to VDH in the past 24 hours.

Scenarios – Transmission Conditions

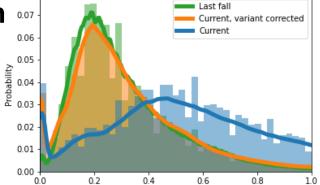
- Variety of factors continue to drive transmission rates
 - Seasonal impact of weather patterns, travel and gatherings, fatigue and premature relaxation of infection control practices
- Waning Immunity: Mean of one year protection (rate of 0.0027) similar to Pfizer study
- Projection Scenarios:
 - Adaptive: Control remains as is currently experienced into the future with assumption that Delta remains as the majority strain
 - Adaptive-FallWinter2020: Starting this week the core drivers of transmission from Sept 2020 – Feb 2021 are coarsely replayed but boosted to account for Delta's increased transmissibility
 - Adaptive-Surge Control: Starting in one week behaviors and mitigation efforts ramp up over a 2-week period culminating in a 25% reduction in transmission



Scenarios – FallWinter2020 Description

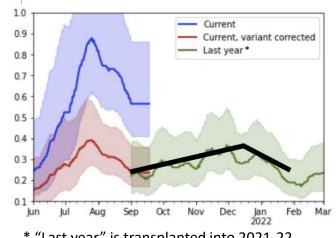
September 2020 – February 2021 saw a strong wave of transmission

- We analyze previous Fall-Winter's wave vs. current Delta driven wave and observe surprising similarities
 - The distribution of fitted model transmissibility is nearly identical between these periods when corrected for Delta's increased transmissibility

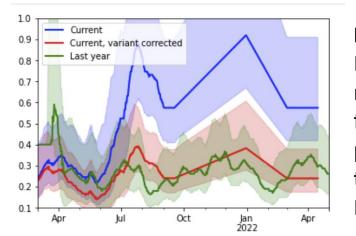


- FallWinter2020 tries to capture the "transmission drivers" from the past and use them as if they were to occur again this season but with Delta variant (compared to ancestral)
 - Use the above analysis of fitted model transmissibilities from Sept 2020 Feb 2021 to guide the future transmissibility from Sept 2021 through Feb 2022, but add the enhanced transmissibility of Delta back in

Fitting: Black line represents the coarsely fitted base transmissibility







Delta enhanced:

Blue trajectory represents current fitted and then projected transmissibility in FallWinter2020

* "Last year" is transplanted into 2021-22

Scenarios – Vaccination Conditions

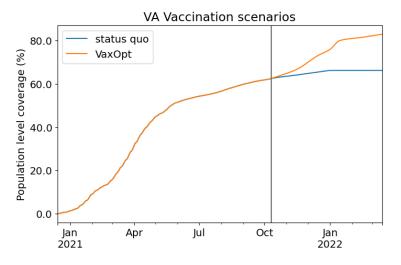
Vaccine Characteristics

- **Pfizer/Moderna**: 50% after first dose, 95% after second dose (3.5 week gap) **J & J**: 67% efficacy after first dose
- Delay to efficacy from doses is 14 days, immunity lasts at least 7m (<u>NEJM</u> study)

Vaccine Administration Scenarios

- Status quo (no label): COVIDcast corrected acceptance estimates (statewide mean is ~80% adults, 65% of population) reached by end of October.
- Optimistic (VaxOpt): Expand VA mean acceptance to include "probably not" (~85% adults) with addition of childhood (5-11 yo) rollout starting in Nov 15th. This follows the same rates as observed of adolescents and results in a net increase of ~10% of population by end of February. Additionally, all counties guaranteed to reach a minimum of 65%, max of 95% by end of December
- Acceptance at county level = regional acceptance +/- relative current vax
- Front-loaded rollout (two-thirds of the remaining in half the time)





	Monthly		Cumu	lative
	status		status	
Date	quo	VaxOpt	quo	VaxOp
2/31/20	110.2K	110.2K	110.2K	110.2K
1/31/21	649.8K	649.8K	760.0K	760.0K
2/28/21	561.7K	561.7K	1.3M	1.3M
3/31/21	1.3M	1.3M	2.6M	2.6M
4/30/21	1.2M	1.2M	3.8M	3.8M
5/31/21	575.8K	575.8K	4.4M	4.4M
6/30/21	243.0K	243.0K	4.6M	4.6M
7/31/21	198.2K	198.2K	4.8M	4.8M
8/31/21	271.6K	271.6K	5.1M	5.1M
9/30/21	177.4K	177.4K	5.3M	5.3M
0/31/21	145.0K	249.9K	5.4M	5.5M
1/30/21	110.5K	432.3K	5.5M	6.0M
2/31/21	122.3K	507.6K	5.7M	6.5M
1/31/22	0	446.2K	5.7M	6.9M
2/28/22	0	103.2K	5.7M	7.0M
3/31/22	0	67.6K	5.7M³2	7.1M

Projection Scenarios – Combined Conditions

Name	Txm Controls	Vax	Description
Adaptive	С	SQ	Likely trajectory based on conditions remaining similar to the current experience
Adaptive-VaxOpt	С	VO	Vaccination through October reaches an optimistically high level of expanded coverage (85%)
Adaptive-SurgeControl	25%	SQ	Transmission rates in the next month reduced through increased control from non-pharmaceutical interventions, with status quo vax and Delta
Adaptive-FallWinter2020	FallWinter 2020	SQ	Transmission rates coarsely follow the rates from last September through this February but are boosted by Delta's enhanced transmissibility
Adaptive-FallWinter2020- VaxOpt	FallWinter 2020	VO	Transmission rates coarsely follow the rates from last September through this February but are boosted by Delta's enhanced transmissibility, with optimistic vax

Transmission Controls: C = Current levels persist into the future

25% = Transmission rates are reduced by 25% with a gradual introduction, concluding in 4 weeks

FallWinter2020 = Transmission rates from Sept 2020 – Feb 2021 are coarsely replayed but boosted by

Delta's increased transmissibility

Vaccinations: SQ = Status quo acceptance leads to low rates of vaccination through the summer

VO = Vaccination acceptance optimistically expands with increased rates through the summer

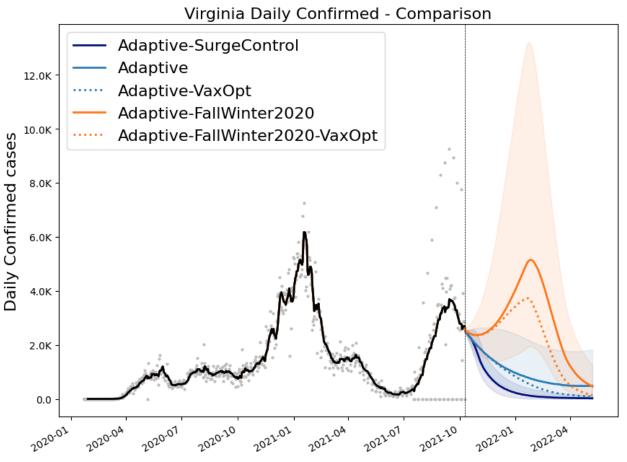
13-Oct-21 33

Model Results

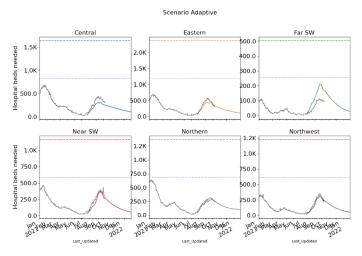


Outcome Projections

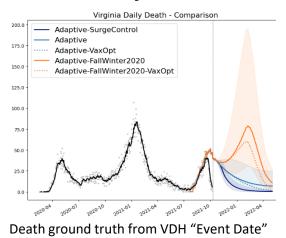
Confirmed cases



Estimated Hospital Occupancy

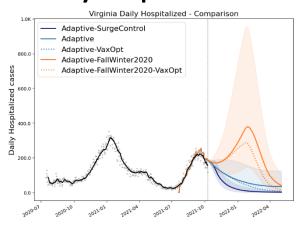


Daily Deaths



data, most recent dates are not complete

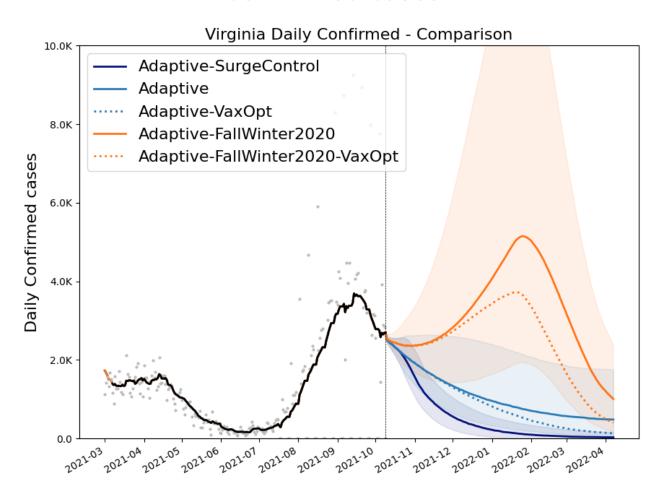
Daily Hospitalized





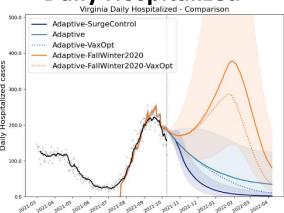
Outcome Projections – Closer Look

Confirmed cases

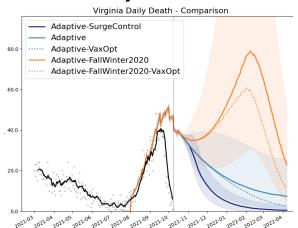




Daily Hospitalized



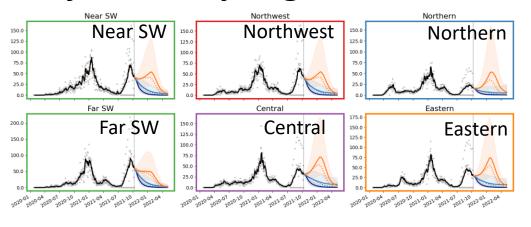
Daily Deaths



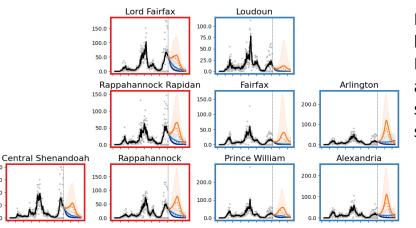
Death ground truth from VDH "Event Date" data, most recent dates are not complete

Detailed Projections: All Scenarios

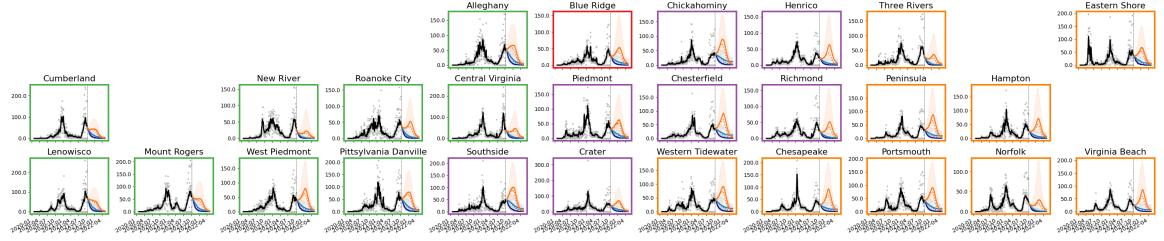
Projections by Region



Projections by District

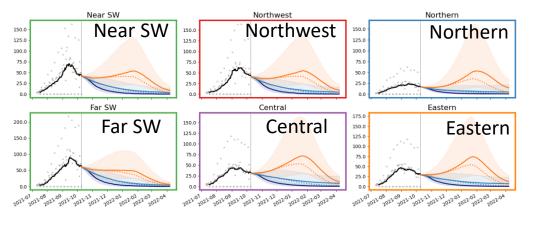


Daily confirmed cases) by rate (per 100K) District (grey with 7-day average in black) with simulation colored by scenario

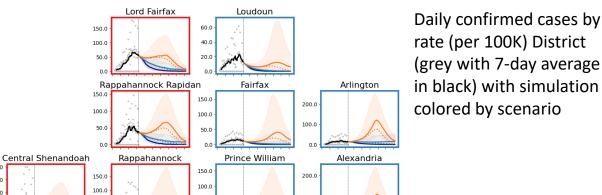


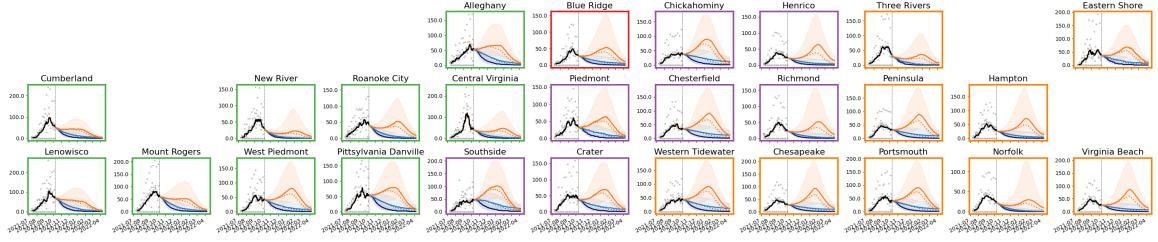
Detailed Projections: All Scenarios - Closer Look

Projections by Region



Projections by District



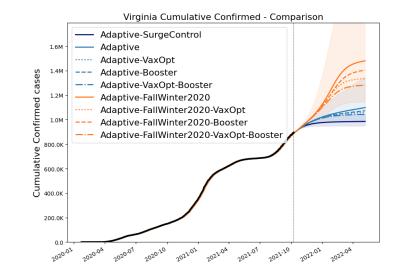


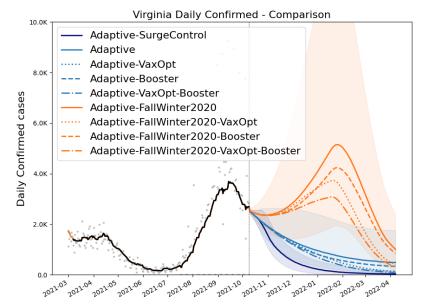
150.0

Impact of Expanded Vaccine Coverage & Boosted Immunity

Vaccination eligibility is expanding (3rd doses, 5-11 year olds, increased acceptance).

- These estimates are preliminary
- 3rd doses: Estimate 40% uptake after 6 months of initial vax, returning waned immunity back to 95% (calibrated to observed additional doses)
- Third doses alone may reduce case counts by ~25K, VaxOpt alone by ~40K, and combined ~55K
- When challenged with another FallWinter wave like 2020, the 3rd dose may reduce cases by ~76K, VaxOpt alone by ~145K, and combined by ~200K
 - Initial analysis of National data suggests moderate initial uptake of 3rd doses, with 30% of doses administered since Sept 26th (3rd dose for larger group authorization)
 - 3M doses over 1st 10 days



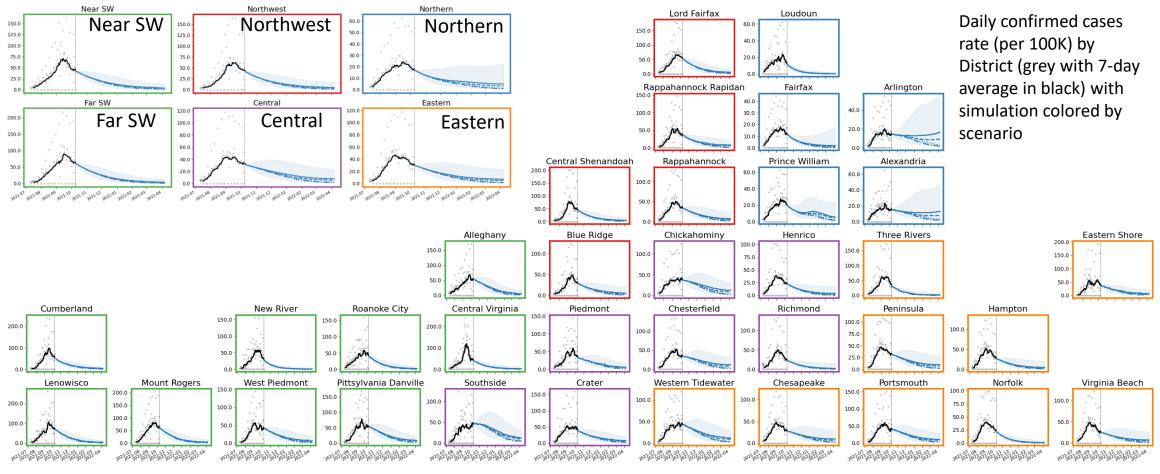




Detailed Projections: Adaptive with Vax Scenarios

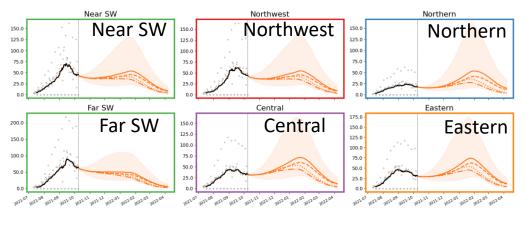
Projections by Region

Projections by District

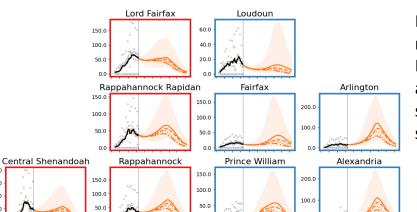


Detailed Projections: Adaptive-FallWinter2020 with Vax Scenarios

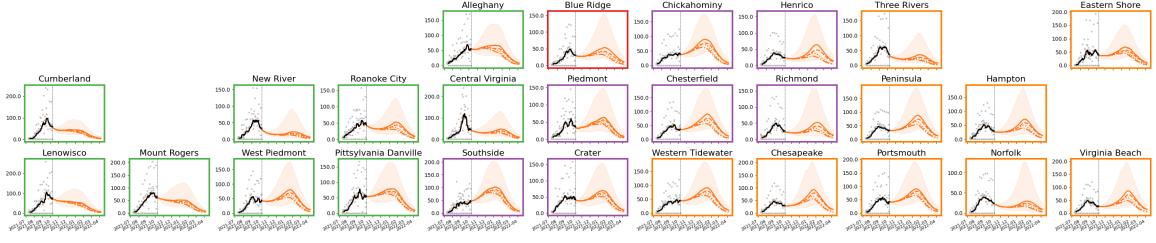
Projections by Region



Projections by District



Daily confirmed cases rate (per 100K) by District (grey with 7-day average in black) with simulation colored by scenario

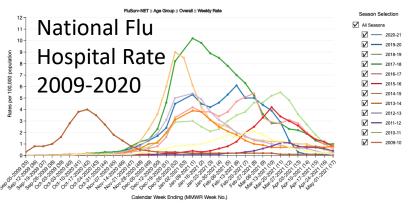


100.0

Impact of Influenza based on Previous Intense Flu Seasons

Augment COVID-19 daily hospitalizations with that of past Influenza seasons

- Include hybrid seasons that use timing of one season but are scaled by severity of another
- Due to limited historical data on Virginia flu hospitalizations currently using national rates applied to VA population



https://gis.cdc.gov/GRASP/Fluview/FluHospRates.html

2009-10 – Pandemic 2009 H1N1 season

2017-18 – Timing and severity of 2017-18 season

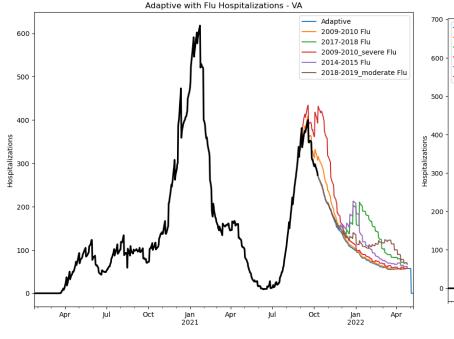
2009-10_severe – Timing of 2009 pandemic (early) with the severity of the 2017-18 season

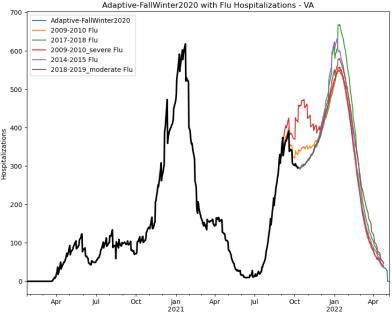
2014-15 – Timing and severity of 2014-15 season

2018-19_moderate – Timing of 2018-19 (late) season with severity of 2014-15



Adaptive – FallWinter2020



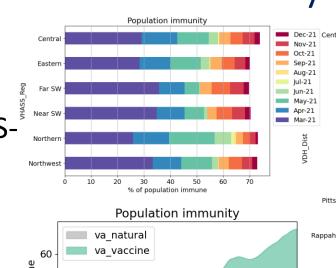


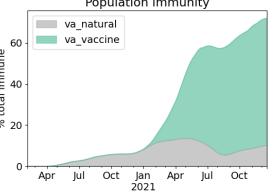


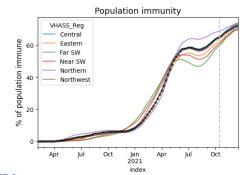
Virginia's Progress on Population Immunity

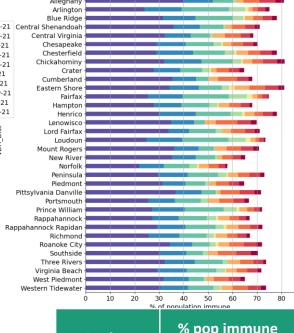
Natural Immunity and Vaccines combine to produce a population level of immunity

- Duration of immunity from infection with SARS-CoV2 still not well understood
 - We assume a conservative 6 month period of protection for these calculations
 - Do **not** factor in variant immune escape
 - Natural immunity is well calibrated to recent seroprevalence surveys
- Vaccine induced immunity is likely to last longer, we assume indefinite protection
 - This also assumes that all administered vaccines remain protective against current and future variants
- Population immunity depends on a very high proportion of the population getting vaccinated
 - Current models track measured seroprevalence









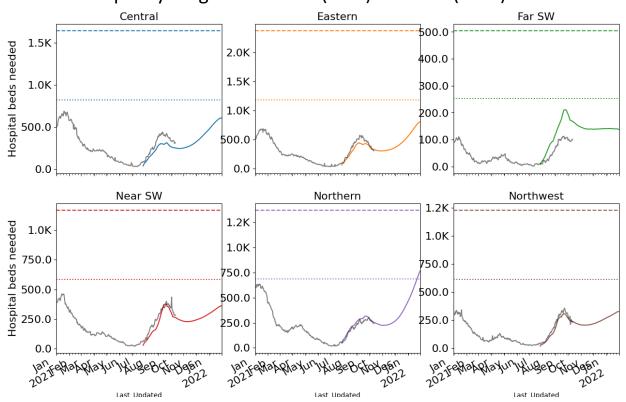
% pop immune (est.)*
65%
61%
59%
60%
68%
64%
64%

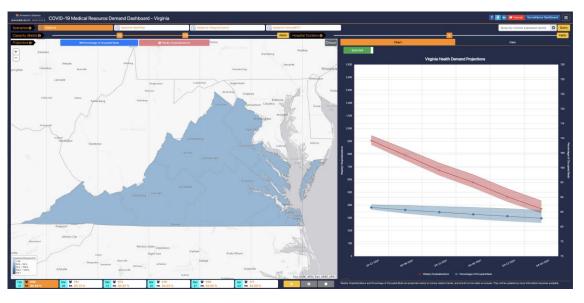
^{*} As of Oct 10, 2021 for entire population

Hospital Demand and Bed Capacity by Region

Capacities* by Region – Adaptive FallWinter2020

COVID-19 capacity ranges from 80% (dots) to 120% (dash) of total beds





https://nssac.bii.virginia.edu/covid-19/vmrddash/

Adaptive FallWinter2020 scenario shows that even with Delta enhanced severity:

- No regions should exceed their surge capacities
- Some regions may exceed initial capacities

* Assumes average length of stay of 8 days



Key Takeaways

Projecting future cases precisely is impossible and unnecessary. Even without perfect projections, we can confidently draw conclusions:

- Case rates in Virginia continue to decline, with nearly all districts declining as well; case rates remain high though the rate of decline has picked up again
- VA 7-day mean daily incidence is slightly down to 29/100K from 32/100K; US is also slightly down to 35/100K (from 39/100K)
- Projections show continued decline across the board, more rapid in some districts than others
- Future case growth remains possible, however, when tested with transmission drivers from last year
- Recent updates:
 - Analysis to show potential impact of Influenza based on past seasons
 - Updated vaccination schedules, adjusted 3rd dose uptake and 5-11 year old timing
 - Adjustment to higher levels of assumed immunity waning (natural and vaccine)

The situation continues to change. Models continue to be updated regularly.



Additional Analyses



Weekly Cases and Hospitalizations

Weekly confirmed cases

Weekly Hospitalizations

Week Ending	Adaptive	Adaptive- VaxOpt	Adaptive- SurgeControl	Adaptive- FallWinter 2020	Adaptive- FallWinter 2020- VaxOpt
10/10/21	18569	18568	18569	18598	18600
10/17/21	16753	16761	16755	17278	17271
10/24/21	15393	15393	15210	16746	16714
10/31/21	14104	14091	12725	16551	16503
11/7/21	12965	12944	9772	16664	16602
11/14/21	11875	11825	7534	17049	16908
11/21/21	10926	10829	5997	17734	17486
11/28/21	10055	9895	4770	18712	18263
12/5/21	9273	9002	3807	19987	19234
12/12/21	8557	8160	3083	21567	20350
12/19/21	7911	7389	2499	23432	21543
12/26/21	7360	6639	2032	25532	22701
1/2/2022	6834	5931	1654	27791	23747
1/9/2022	6372	5267	1359	30288	24770

Week Ending	Adaptive	Adaptive- VaxOpt	Adaptive- SurgeControl	Adaptive- FallWinter 2020	Adaptive- FallWinter 2020- VaxOpt
10/10/21	1326	1326	1326	1327	1327
10/17/21	1245	1245	1245	1264	1264
10/24/21	1148	1148	1146	1218	1217
10/31/21	1054	1052	1001	1195	1191
11/7/21	971	969	794	1198	1192
11/14/21	892	888	607	1221	1211
11/21/21	822	816	480	1263	1248
11/28/21	758	749	382	1329	1303
12/5/21	699	684	305	1414	1371
12/12/21	646	623	245	1521	1456
12/19/21	596	568	199	1651	1550
12/26/21	554	513	162	1799	1646
1/2/2022	513	463	131	1960	1740
1/9/2022	478	414	107	2136	1831

Overview of relevant on-going studies

Other projects coordinated with CDC and VDH:

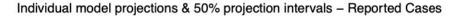
- Scenario Modeling Hub: Consortium of academic teams coordinated via MIDAS / CDC to that provides regular national projections based on timely scenarios
- Genomic Surveillance: Analyses of genomic sequencing data, VA surveillance data, and collaboration with VA DCLS to identify sample sizes needed to detect and track outbreaks driven by introduction of new variants etc.
- Mobility Data driven Mobile Vaccine Clinic Site Selection: Collaboration with VDH state and local, Stanford, and SafeGraph to leverage anonymized cell data to help identify

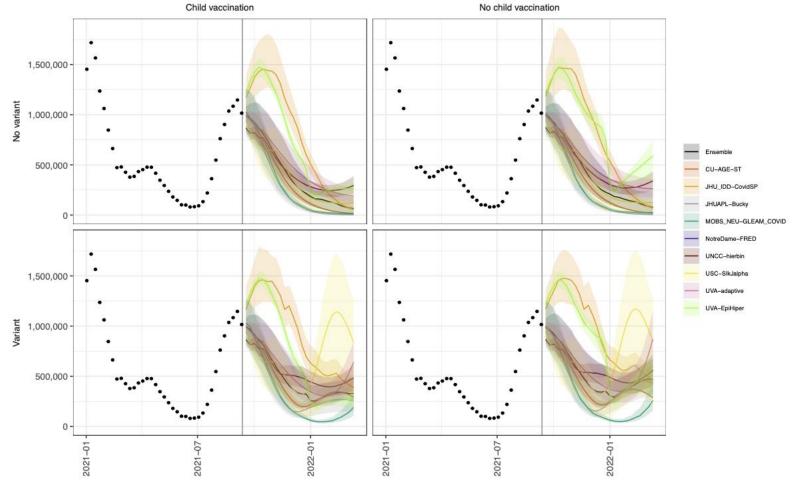
COVID-19 Scenario Modeling Hub

Collaboration of multiple academic teams to provide national and state-by-state level projections for 4 aligned scenarios that vary vaccine rates (high – low) and impact of the Delta variant (high and low)

- Round 9 released to assist in support of 5-11 vax consideration (ACIP meeting Sept 22-23)
- Rounds 4-8 now available Round 4 Results were published May 5th, 2021 in MMWR

https://covid19scenariomodelinghub.org/viz.html





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COVID-19 Scenario Modeling Hub – Round 7

Round 7 scenarios explore the effects of a variant similar to Delta (B.1.617.2) against different backgrounds of vaccination. Includes some vax escape

Vaccinations in 5-11 start in Nov

Follows same rates as adolescents

Emerging Variant Impact (5% prevalence on Nov 15)

• 50% boost as it eventually predominates

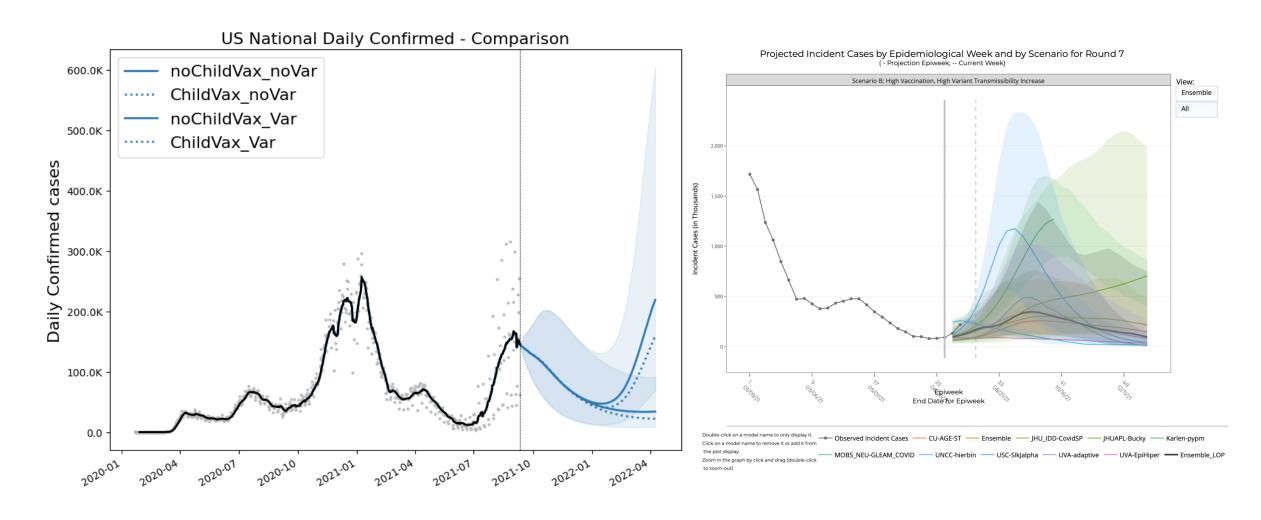
We consider a 2x2 scenario design, where childhood vaccination (5-11 years) is on the first axis, and a change in virus transmissibility is on the second axis. The second axis reflects a stress test, illustrating the potential impact of a new variant arising during the projection period:

	The same mix of variants circulate throughout the projection period. No change in virus transmissibility.	A more transmissible variant emerges, comprising 1% of circulating viruses on Nov 15 . The new variant is 1.5X as transmissible as viruses circulating at the beginning of the projection period.
Vaccination among 5-11yrs is approved and immunization begins on Nov 1. Each state's uptake rate reflects the percent coverage increases observed for 12-17-year-olds since distribution began on May 13.	A	С
No vaccination for children under 12	В	D

https://covid19scenariomodelinghub.org/viz.html

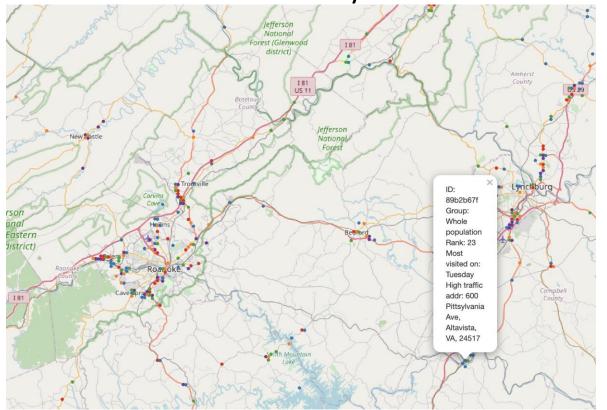
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Modeling Hub – Round 9 Prelim Results



Data Recommended Mobile Vax Clinic Sites

Detailed and Timely Locations



Data Delivered and Disseminated to Locals

Provides a list of areas most visited by a given demographic group based on SafeGraph mobility data that links visits to specific sites and the home Census Block Group of the anonymized visitors

Demographic Groups: Black, Lantinx, Young Adults (20-40), Unvaccinated, and Whole Population

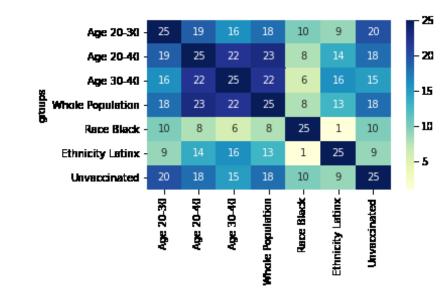
Data Included: Rank, Weight, most visited Day of Week, Highly Visited Address, and Lat-Long of area

Goal: Provide frequently visited locations based on populations and vaccination levels one desires to reach **Example:** List of location in the Southside frequented by 20-40 year olds



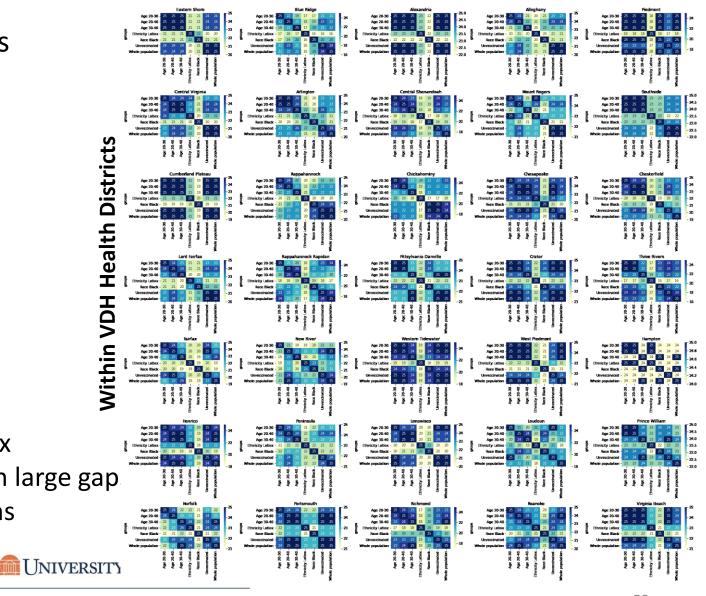
Data Recommended Mobile Vax Clinic Sites

Overlap of locations between groups



Different groups visit different areas

- Least overlap between Black and Latinx
- Overlap in ages highest, but drops with large gap
- Districts have different overlap patterns



State Level

Estimating Daily Reproductive Number

October 11th Estimates

Region	Date Confirmed R _e	Date Confirmed Diff Last Week
State-wide	0.434	-0.581
Central	0.491	-0.550

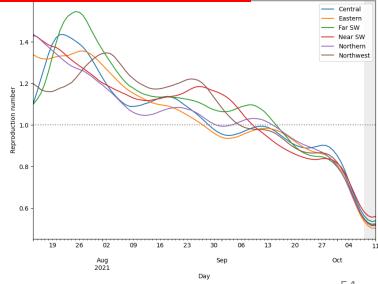


Near SVv	0.492	-0.491
Northern	0.462	-0.527
Northwest	0.456	-0.564

Methodology

- Wallinga-Teunis method (EpiEstim¹) for cases by **confirmation date**
- Serial interval: updated to discrete distribution from observations (mean=4.3, Flaxman et al, Nature 2020)
- Using Confirmation date since due to increasingly unstable estimates from onset date due to backfill

1. Anne Cori, Neil M. Ferguson, Christophe Fraser, Simon Cauchemez. A New Framework and Software to Estimate Time-Varying Reproduction Numbers During Epidemics. American Journal of Epidemiology, Volume 178, Issue 9, 1 November 2013, Pages 1505–1512, https://doi.org/10.1093/aje/kwt133



References

Venkatramanan, S., et al. "Optimizing spatial allocation of seasonal influenza vaccine under temporal constraints." *PLoS Computational Biology* 15.9 (2019): e1007111.

Arindam Fadikar, Dave Higdon, Jiangzhuo Chen, Bryan Lewis, Srinivasan Venkatramanan, and Madhav Marathe. Calibrating a stochastic, agent-based model using quantile-based emulation. SIAM/ASA Journal on Uncertainty Quantification, 6(4):1685–1706, 2018.

Adiga, Aniruddha, Srinivasan Venkatramanan, Akhil Peddireddy, et al. "Evaluating the impact of international airline suspensions on COVID-19 direct importation risk." *medRxiv* (2020)

NSSAC. PatchSim: Code for simulating the metapopulation SEIR model. https://github.com/NSSAC/PatchSim

Virginia Department of Health. COVID-19 in Virginia. http://www.vdh.virginia.gov/coronavirus/

Biocomplexity Institute. COVID-19 Surveillance Dashboard. https://nssac.bii.virginia.edu/covid-19/dashboard/

Google. COVID-19 community mobility reports. https://www.google.com/covid19/mobility/

Biocomplexity page for data and other resources related to COVID-19: https://covid19.biocomplexity.virginia.edu/



Questions?

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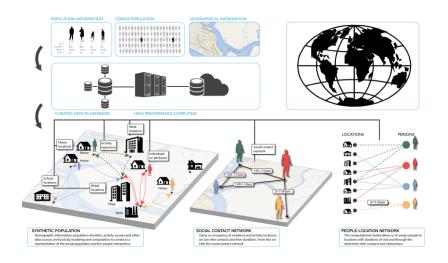
Supplemental Slides



Agent-based Model (ABM)

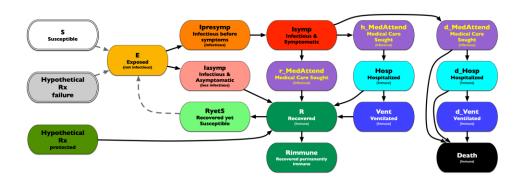
EpiHiper: Distributed network-based stochastic disease transmission simulations

- Assess the impact on transmission under different conditions
- Assess the impacts of contact tracing



Synthetic Population

- Census derived age and household structure
- Time-Use survey driven activities at appropriate locations



Detailed Disease Course of COVID-19

- Literature based probabilities of outcomes with appropriate delays
- Varying levels of infectiousness
- Hypothetical treatments for future developments

